(FILE 'HOME' ENTERED AT 10:36:32 ON 23 MAR 2002)

INDEX 'ADISALERTS, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA,

CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 10:36:57 ON 23 MAR 2002

## SEA NISA

```
_____
   FILE AGRICOLA
45
15
     FILE AQUASCI
     FILE BIOBUSINESS
12
56
     FILE BIOSIS
    FILE BIOTECHABS
21
     FILE BIOTECHDS
21
     FILE BIOTECHNO
36
    FILE CABA
52
90
    FILE CAPLUS
     FILE CEABA-VTB
7
     FILE CIN
2
     FILE CONFSCI
1
     FILE DGENE
50
     FILE DRUGLAUNCH
1
1
     FILE DRUGMONOG2
45
     FILE EMBASE
27
    FILE ESBIOBASE
54
    FILE FOMAD
    FILE FROSTI
11
    FILE FSTA
38
4
    FILE GENBANK
    FILE HEALSAFE
2
    FILE IFIPAT
2
    FILE JICST-EPLUS
14
    FILE LIFESCI
40
62
    FILE MEDLINE
```

6 FILE WPIDS 6 FILE WPINDEX QUE NISA

------

38

3

61

478 81

16

23

L1

L2

L3

1

FILE 'PROMT, CAPLUS, SCISEARCH, MEDLINE, BIOSIS, PASCAL, AGRICOLA' ENTERED AT 10:38:11 ON 23 MAR 2002

7 S L1(S)BETA-GALACTOSIDASE

FILE NTIS

FILE OCEAN

FILE PROMT

FILE SCISEARCH FILE TOXCENTER

FILE USPATFULL

FILE PASCAL FILE PHIN

2 DUP REM L2 (5 DUPLICATES REMOVED)

L3 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:284076 CAPLUS

DOCUMENT NUMBER: 134:309806

TITLE: Lactose hydrolysis using recombinant lactic acid

bacteria producing high levels of

.beta.-galactosidase

INVENTOR (S):

Ruch, Frank E.

PATENT ASSIGNEE(S):

Protein Scientific, Inc., USA

SOURCE:

PCT Int. Appl., 51 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

```
PATENT NO.
                   KIND DATE
                                        APPLICATION NO. DATE
                                         WO 2001027247 A2 20010419
WO 2001027247 A3 20011018
                                        WO 2000-US41121 20001006
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
            CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                      US 1999-158668P P 19991008
                                      US 2000-542121 A1 20000404
```

AB The invention features methods and compns. for rapidly and effectively hydrolyzing lactose using recombinant lactic acid bacteria that produce high levels of .beta.-galactosidase, permeabilizing the bacteria such that

lactose can enter the cell and be hydrolyzed by the highly concd. .beta.-galactosidase contained herein. The invention further features a reduced lactose diary product, e.g., milk. The invention features also lactase microcarriers as an oral prophylactic against the clin. condition of lactose intolerance.

L3 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 1

ACCESSION NUMBER:

1998:811148 CAPLUS

DOCUMENT NUMBER:

130:205817

TITLE:

Nisin independent induction of the nisA promoter in Lactococcus lactis during growth in lactose or

galactose

AUTHOR (S):

Chandrapati, Sailaja; O'Sullivan, Daniel J.

CORPORATE SOURCE:

Department of Food Science and Nutrition, University

of Minnesota, St. Paul, MN, 55108, USA

SOURCE:

FEMS Microbiol. Lett. (1999), 170(1), 191-198

CODEN: FMLED7; ISSN: 0378-1097

PUBLISHER:

Elsevier Science B.V.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB Nisin biosynthesis is autoregulated extracellularly by the mature and modified peptide. To investigate other regulatory effects on nisin biosynthesis, a transcription fusion of the nisA promoter from Lactococcus

lactis ATCC 11454 to the promoterless lacZ gene from Streptococcus

thermophilus was constructed. This fusion construct, pDOC99, expressed .beta.-galactoside in L. lactis ATCC 11454 growing in M17 medium contg. glucose (M17G). Consistent with the known model for transcription of nisA, pDOC99 did not express .beta.-galactosidase in the non-nisin producer, L. lactis LM0230 grown in M17G, unless the nisRK genes (cloned in pDOC23) were included in trans

and

nisin was added to the medium. Growth of this strain in M17 contg. lactose or galactose, resulted in nisA transcription, even in the absence of exogenous nisin. This expression was independent of pDOC23. Furthermore, nisA transcription in L. lactis LM0230(pDOC99) grown in M17G could be induced by the addn. of exogenous galactose, with max. induction occurring at concns. 5 mM.

REFERENCE COUNT:

14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR

THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 18 OF 19 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 13

ACCESSION NUMBER: 1988:568937 CAPLUS

DOCUMENT NUMBER: 109:168937

Fermentation of lactose by Zymomonas mobilis carrying TITLE:

a Lac+ recombinant plasmid

AUTHOR(S): Yanase, Hideshi; Kurii, Junn; Tonomura, Kenzo

Coll. Agric., Univ. Osaka Prefect., Osaka, 591, Japan CORPORATE SOURCE:

J. Ferment. Technol. (1988), 66(4), 409-15 SOURCE:

CODEN: JFTED8; ISSN: 0385-6380

DOCUMENT TYPE: Journal LANGUAGE: English

Lac+ recombinant plasmids encoding a .beta. -

galactosidase fused protein and lactose permease of Escherichia coli were introduced into Z. mobilis. The fused protein was expressed

with 450 to 5860 Miller units of .beta.-

galactosidase activity, and functioned as lactase. Raffinose uptake by Z. mobilis CP4 was enhanced in the plasmid-carrying strain over the plasmid-free strain, suggesting that the lactose permease was functioning in the organism. Z. mobilis Carrying the plasmid could

produce EtOH from lactose and whey, but could not grown on lactose as sole

C source. Its growth was inhibited by either galactose or the galactose liberated from lactose.

L9 ANSWER 15 OF 19 FSTA COPYRIGHT 2002 IFIS

ACCESSION NUMBER: 1997(02):B0111 FST

TITLE: Characterization of an oxygen-dependent inducible

promoter system, the nar promoter, and Escherichia

coli with an inactivated nar operon.

AUTHOR: Jintae Lee; Moo Hwan Cho; Jongwon Lee

CORPORATE SOURCE: Correspondence (Reprint) address, Jongwon Lee, Dep.

of

gene

Biochem., Sch. of Med., Catholic Univ. of

Taegu-Hyosung, 3056-6, Daemyung 4-Dong, Nam-Gu, Taegu

705-034, Korea. Tel. 82-53-650-4471. Fax

82-53-621-4106

SOURCE: Biotechnology and Bioengineering, (1996) 52 (5)

572-578, 22 ref. ISSN: 0006-3592

DOCUMENT TYPE: Journal LANGUAGE: English

AB The nar promoter of Escherichia coli, which is optimally induced in the presence of nitrate under anaerobic conditions, was characterized in order

to ascertain its usefulness as an inducible promoter. The nar promoter was

expressed in an E. coli strain having a mutant nar operon which does not express active nitrate reductase. A plasmid containing the lacZ gene, expressing .beta.-galactosidase, instead of the structural genes of the nar operon was used to assay induction of the nar promoter. Optimal conditions for nar induction were analysed. Results showed that induction of the nar promoter was optimal when E. coli was grown initially in the presence of 1% nitrate. Expression of the lacZ

was not affected by molybdate ions. The amount of .beta.galactosidase per cell and per medium vol. was max. when E. coli
was grown under aerobic conditions to an optical density (at 600 nm) of
1.7; induction of the nar promoter was observed by lowering dissolved
0.sub.2 concn. to microanaerobic levels (1-2%). After approx. 6 h
induction, specific .beta.-galactosidase activity was
36 000 Miller units, equivalent to 35% of total
cellular proteins, which was confirmed by SDS-PAGE. The specific activity
of .beta.-galactosidase expressed from the nar

L9 ANSWER 14 OF 19 FSTA COPYRIGHT 2002 IFIS ACCESSION NUMBER: 1996(06):B0141 FSTA

TITLE: High-level expression of lacZ under control of the

tac

or trp promoter using runaway replication vectors in

Escherichia coli.

AUTHOR: Kidwell, J.; Kolibachuk, D.; Dennis, D.

CORPORATE SOURCE: Correspondence (Reprint) address, D. Dennis, Dep. of

Biol., James Madison Univ., Harrisonburg, VA 22807,

USA

SOURCE: Biotechnology and Bioengineering, (1996) 50 (1)

108-114, 24 ref. ISSN: 0006-3592

DOCUMENT TYPE: Journal LANGUAGE: English

AB The Escherichia coli lacZ gene, encoding .beta.-

galactosidase, was placed under control of the trp or tac promoter
in the runaway replication vectors pRA95 and pRA96, in which copy number
is thermally regulated. Expression of lacZ was examined in transformed

cells containing these plasmids. Increasing the temp. increased

expression

of the lacZ gene; 41.degree.C was the optimum temp. for thermal induction of gene expression. Induction of gene expression using isopropyl-.beta.-D-thiogalactopyranoside (IPTG) or 3-.beta.-indoleacrylic acid IAA did not significantly enhance thermal induction of gene expression. In thermally induced strains harbouring the tac promoter, a lag period of approx. 1.5 h was observed prior to .beta.-galactosidase production; no apparent lag was observed in strains possessing the trp promoter. Max. .beta.-galactosidase

levels (up to 46 000 **Miller units**) were produced using

a trp promoter on pRA96, having a basal copy number of 10; enzyme levels

L9 ANSWER 12 OF 19 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 11

ACCESSION NUMBER: 1996:175149 CAPLUS

DOCUMENT NUMBER: 124:222089

TITLE: High-level expression of lacZ under control of the

tac

or trp promoter using runaway replication vectors in

Escherichia coli

AUTHOR(S): Kidwell, John; Kolibachuk; Dennis, Douglas

CORPORATE SOURCE: Dep. Biol., James Madison Univ., Harrisonburg, VA,

22807, USA

SOURCE: Biotechnol. Bioeng. (1996), 50(1), 108-14

CODEN: BIBIAU; ISSN: 0006-3592

DOCUMENT TYPE: Journal LANGUAGE: English

AB To det. the utility of coupling runaway replication to the expression of cloned genes under the control of strong promoters, lacZ transcriptional fusions to the trp or tac promoter (Ptrp or Ptac) were constructed using plasmids in which the copy no. is thermally regulated. Cells contg.

these

plasmids were able to produce .beta.-galactosidase to levels between 3700 and 46,000 Miller units when induced only by a temp. upshift. The addn. of the appropriate chem. inducer, either IPTG (isopropyl-.beta.-D-thiogalactopyranoside) or IAA (3-.beta.-indoleacrylic acid), did not significantly enhance the thermal induction. The Ptac-controlled and Ptrp-controlled lacZ induction differed slightly in that the Ptac-controlled thermal induction exhibited a lag of approx. 1.5 h as compared to both chem. and thermal induction, whereas in the case of Ptrp-controlled induction, an increase in . beta.-galactosidase expression above background occurred at approx. the same time regardless of the means of induction. The best vector, a Ptrp-controlled lacZ fusion carried on a runaway replication vector having a basal copy no. of 10, was able to mediate the expression of .beta.-galactosidase to approx. 40,000

## Miller units of .beta.-galactosidase

comprising 25% of the total cell protein at 17 h postinduction under optimal conditions for protein yield. In these cells, lysis occurred as lacZ was maximally expressed. Under noninducing conditions, the plasmids were stable for at least 60 generations in the absence of antibiotic in

L9 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 7

ACCESSION NUMBER: 1998:803860 CAPLUS

DOCUMENT NUMBER: 130:205625

TITLE: Development of a plasmid vector for overproduction of

.beta.-galactosidase in

Escherichia coli by using genetic components of groEx

from symbiotic bacteria in Amoeba proteus Lee, Jung Eun; Ahn, Eun Young; Ahn, Tae In

Department of Biology Education, Seoul National University, Seoul, 151-742, S. Korea

J. Microbiol. Biotechnol. (1998), 8(5), 509-516

CODEN: JOMBES; ISSN: 1017-7825

PUBLISHER: Korean Society for Applied Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

AUTHOR (S):

SOURCE:

CORPORATE SOURCE:

AB A plasmid vector, pXGPRMATG-lac-Tgx, was developed for overprodn. of .

beta.-galactosidase in Escherichia coli using the
genetic components of groEx, a heat-shock gene cloned from symbiotic
X-bacteria in Amoeba proteus. The vector is composed of intragenic
promoters P3 and P4 of groEx, the structural gene of lac operon,
transcription terminator signals of lac and groEx, and ColE1 and amp' of
pBluescript SKII. The optimized host, E. coli DH5.alpha., transformed
with the vector constitutively produced 117,310-171,961 Miller
units of .beta.-galactosidase per mg protein
in crude ext. The amt. of enzyme in crude ext. was 53% of total
water-sol. proteins. About 43% of the enzyme could be purified to a
specific activity of 322,249 Miller units/mg protein
after two-fold purifn., using two cycles of pptn. with ammonium sulfate
and one step of gel filtration. Thus, the expression system developed in

this study presents a low cost and simple method for purifying

L3 ANSWER 38 OF 45 SCISEARCH COPYRIGHT 2002 ISI (R)

ACCESSION NUMBER: 91:688106 SCISEARCH

THE GENUINE ARTICLE: GV187

TITLE: THE BACTERIOCIN LACTOCOCCIN-A SPECIFICALLY INCREASES

PERMEABILITY OF LACTOCOCCAL CYTOPLASMIC MEMBRANES
IN A VOLTAGE-INDEPENDENT, PROTEIN-MEDIATED MANNER

AUTHOR: VANBELKUM M J (Reprint); KOK J; VENEMA G; HOLO H; NES I

F;

KONINGS W N; ABEE T

CORPORATE SOURCE: UNIV GRONINGEN, DEPT GENET, KERKLAAN 80, 9751 NN HAREN,

NETHERLANDS (Reprint); NLVF, MICROBIAL GENE TECHNOL LAB, N-1432 AS, NORWAY; UNIV GRONINGEN, DEPT MICROBIOL, 9751

NN

HAREN, NETHERLANDS

COUNTRY OF AUTHOR: NETHERLANDS; NORWAY

SOURCE: JOURNAL OF BACTERIO

JOURNAL OF BACTERIOLOGY, (1991) Vol. 173, No. 24, pp.

7934-7941.

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE LANGUAGE: ENGLISH

REFERENCE COUNT: 36

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

Lactococcin A is a bacteriocin produced by Lactococcus lactis. Its structural gene has recently been cloned and sequenced (M. J. van Belkum. B. J. Hayema, R. E. Jeeninga, J. Kok, and G. Venema, Appl. Environ. Microbiol. 57:492-498, 1991). Purified lactococcin A increased the permeability of the cytoplasmic membrane of L. lactis and dissipated the membrane potential. A significantly higher concentration of lactococcin A was needed to dissipate the membrane potential in an immune strain of L. lactis. Lactococcin A at low concentrations (0.029-mu-g/mg of protein) inhibited secondary and phosphate-bond driven transport of amino acids in sensitive cells and caused efflux of preaccumulated amino acids. Accumulation of amino acids by immune cells was not affected by this concentration of lactococcin A. Lactococcin A also inhibited proton motive force-driven leucine uptake

and

leucine counterflow in membrane vesicles of the sensitive strain but not in membrane vesicles of the immune strain. These observations indicate that lactococcin A makes the membrane permeable for leucine in the presence or absence of a proton motive force and that the immunity factor(s) is membrane linked. Membrane vesicles of Clostridium acetobutylicum, Bacillus subtilis, and Escherichia coli were not affected by lactococcin A, nor were liposomes derived from phospholipids of L. lactis. These results indicate that lactococcin A acts on the cytoplasmic membrane and is very specific towards lactococci. The combined results obtained with cells, vesicles, and liposomes suggest that the specificity of lactococcin A may be mediated by a receptor protein associated with

the

cytoplasmic membra